## **Claims**

- 1.-39. (canceled)
- 40. (currently amended) A method of inducing senescence of a cell, comprising altering the level of a nucleostemin polypeptide comprising an amino acid sequence at least 80%-95% identical to SEQ ID NO: [[6]10, wherein the amino acid sequence has the biological and functional characteristics of the nucleostemin polypeptide set forth as SEQ ID NO: 10, wherein altering the level of the nucleostemin polypeptide comprises:

  contacting the cell with an agent that reduces or increases the level of nucleostemin; and

<u>determining senescence of the cell</u>, thereby inducing senescence of the cell.

- 41. (withdrawn and currently amended) The method of claim 40, wherein altering the level of the nucleostemin polypeptide comprises increasing the level of the <u>nucleostemin</u> polypeptide.
- 42. (currently amended) The method of claim 40, wherein altering the level of the nucleostemin polypeptide comprises decreasing the level of the <u>nucleostemin</u> polypeptide.
- 43. (original) The method of claim 40, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 6.
- 44. (original) The method of claim 40, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 4.
- 45. (original) The method of claim 40, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 2.
- 46. (original) The method of claim 40, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 10.
  - 47. (original) The method of claim 40, wherein the cell is a tumor cell.

- 48. (currently amended) The method of <u>clam-claim</u> 40, wherein the cell is a stem cell.
  - 49. (original) The method of claim 40, wherein the cell is *in vitro*.
  - 50. (original) The method of claim 40, wherein the cell is *in vivo*.
- 51. (currently amended) A method of inducing senescence of a tumor cell in a subject, comprising

administering to the subject a therapeutically effective amount of an agent that <u>identified</u> as one that alters the level of a nucleostemin polypeptide comprising an amino acid sequence at least 80%-95% identical to SEQ ID NO: 6, wherein the amino acid sequence has the biological and functional characteristics of the nucleostemin polypeptide set forth as SEQ ID NO: 6; and

measuring the senescence of the tumor cell in the subject, thereby inducing senescence of the tumor cell in the subject.

- 52. (original) The method of claim 51, wherein the agent is a small inhibitory RNA that specifically binds a polynucleotide encoding the nucleostemin polypeptide.
- 53. (withdrawn) The method of claim 51, wherein the agent is a polynucleotide encoding a nucleostemin polypeptide.
  - 54. (withdrawn) The method of claim 51, wherein the agent is a p53.
  - 55. (canceled)
- 56. (currently amended) The method of claim [[46]]48, wherein the <u>stem</u> cell is a central nervous system (CNS) stem <del>tumor</del> cell.
  - 57.-60. (canceled)

61. (currently amended) The method of claim [[46]]40, wherein altering the level of the nucleostemin polypeptide comprises introducing into the cell a small inhibitory RNA that specifically binds a polynucleotide encoding the nucleostemin polypeptide.

## 62.-63. (cancelled)

- 64. (withdrawn) The method of claim 40, wherein altering the level of the nucleostemin polypeptide comprises increasing transcription of a nucleic acid sequence encoding the nucleostemin polypeptide.
- 65. (withdrawn) The method of claim 40, wherein altering the level of the nucleostemin polypeptide comprises altering the amount of the polypeptide bound to p53.
- 66. (currently amended) The method of claim [[40]]61, wherein altering the level of the nucleostemin polypeptide comprises introducing into the cell a the small inhibitory RNA consists of a nucleic acid sequence as set forth as SEQ ID NO: 7 that specifically binds a polynucleotide encoding the nucleostemin polypeptide.
- 67. (new) The method of claim 52, wherein the small inhibitory RNA consists of a nucleic acid sequence as set forth as SEQ ID NO: 7.
- 68. (new) A method of inducing senescence of a stem cell, comprising reducing the level of a nucleostemin polypeptide comprising an amino acid sequence at least 95% identical to SEQ ID NO: 10, wherein the amino acid sequence has the biological and functional characteristics of the nucleostemin polypeptide set forth as SEQ ID NO: 6, and wherein reducing the level of the nucleostemin polypeptide comprises:

contacting the stem cell with a small inhibitory RNA identified as one that reduces the level of nucleostemin; and

evaluating the cell for senescence, thereby inducing senescence of the cell.

- 69. (new) The method of claim 68, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 6.
- 70. (new) The method of claim 68, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 4.
- 71. (new) The method of claim 68, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 2.
- 72. (new) The method of claim 68, wherein the nucleostemin polypeptide comprises an amino acid sequence set forth as SEQ ID NO: 10.
  - 73. (new) The method of claim 68, wherein the stem cell is a CNS stem cell.
- 74. (new) The method of claim 68, wherein the small inhibitory RNA consists of a nucleic acid sequence as set forth as SEQ ID NO: 7.
  - 75. (new) The method of claim 47, wherein the tumor cell is an osteocarcinoma cell.
  - 76. (new) The method of claim 75, wherein the osteocarcinoma cell is an U2OS cell.